DEPARTMENT OF HEALTH AND HUMAN SERVICES

Food and Drug Administration

[Docket Nos. FDA-2020-P-2244 and FDA-2020-P-2245]

Determination That ISOPTIN (Verapamil Hydrochloride) Tablets 40 Milligrams, 80 Milligrams, and 120 Milligrams, and CALAN (Verapamil Hydrochloride) Tablets, 40 Milligrams, 80 Milligrams, 120 Milligrams, and 160 Milligrams, Were Not Withdrawn from Sale for Reasons of Safety or Effectiveness

AGENCY: Food and Drug Administration, HHS.

ACTION: Notice.

SUMMARY: The Food and Drug Administration (FDA, Agency, or we) has determined that ISOPTIN (verapamil hydrochloride) tablets, 40 milligrams (mg), 80 mg, and 120 mg, and CALAN (verapamil hydrochloride) tablets, 40 mg, 80 mg, 120 mg, and 160 mg, were not withdrawn from sale for reasons of safety or effectiveness. This determination will allow FDA to approve abbreviated new drug applications (ANDAs) for ISOPTIN (verapamil hydrochloride) tablets, 40 mg, 80 mg, and 120 mg, and CALAN (verapamil hydrochloride) tablets, 40 mg, 80 mg, 120 mg, and 160 mg, if all other legal and regulatory requirements are met.

FOR FURTHER INFORMATION CONTACT: Nikki Mueller, Center for Drug Evaluation and Research, Food and Drug Administration, 10903 New Hampshire Ave., Bldg. 51, Rm. 6280, Silver Spring, MD  20993-0002, 301-796-3601, Nicole.Mueller@fda.hhs.gov.

SUPPLEMENTARY INFORMATION: In 1984, Congress enacted the Drug Price Competition and Patent Term Restoration Act of 1984 (Pub. L. 98-417) (the 1984 amendments), which authorized the approval of duplicate versions of drug products under an ANDA procedure. ANDA applicants must, with certain exceptions, show that the drug for which they are seeking approval contains the same active ingredient in the same strength and dosage form as the “listed drug,” which is a version of the drug that was previously approved. ANDA applicants do not
have to repeat the extensive clinical testing otherwise necessary to gain approval of a new drug application (NDA).

The 1984 amendments include what is now section 505(j)(7) of the Federal Food, Drug, and Cosmetic Act (21 U.S.C. 355(j)(7)), which requires FDA to publish a list of all approved drugs. FDA publishes this list as part of the “Approved Drug Products With Therapeutic Equivalence Evaluations,” which is known generally as the “Orange Book.” Under FDA regulations, drugs are removed from the list if the Agency withdraws or suspends approval of the drug’s NDA or ANDA for reasons of safety or effectiveness or if FDA determines that the listed drug was withdrawn from sale for reasons of safety or effectiveness (21 CFR 314.162).

A person may petition the Agency to determine, or the Agency may determine on its own initiative, whether a listed drug was withdrawn from sale for reasons of safety or effectiveness. This determination may be made at any time after the drug has been withdrawn from sale, but must be made prior to approving an ANDA that refers to the listed drug (§ 314.161 (21 CFR 314.161)). FDA may not approve an ANDA that does not refer to a listed drug.

ISOPTIN (verapamil hydrochloride) tablets, 40 mg, 80 mg, and 120 mg, are the subject of NDA 018593, held by Mt. Adams Technologies, LLC, and initially approved on March 8, 1982. CALAN (verapamil hydrochloride) tablets, 40 mg, 80 mg, 120 mg, and 160 mg, are the subject of NDA 018817, held by Pfizer Inc., and initially approved on September 10, 1984. ISOPTIN and CALAN are indicated for the treatment of angina, arrhythmias, and essential hypertension.

Center Laboratories, Inc., submitted two citizen petitions dated November 30, 2020 (Docket Nos. FDA-2020-P-2244 and FDA-2020-P-2245), under 21 CFR 10.30, requesting that the Agency determine whether ISOPTIN (verapamil hydrochloride) tablets, 40 mg, 80 mg, and 120 mg, and CALAN (verapamil hydrochloride) tablets, 40 mg, 80 mg, 120 mg, and 160 mg, were withdrawn from sale for reasons of safety or effectiveness.
After considering the citizen petitions and reviewing Agency records and based on the information we have at this time, FDA has determined under § 314.161 that ISOPTIN (verapamil hydrochloride) tablets, 40 mg, 80 mg, and 120 mg, and CALAN (verapamil hydrochloride) tablets, 40 mg, 80 mg, 120 mg, and 160 mg, were not withdrawn for reasons of safety or effectiveness. The petitioner has identified no data or other information suggesting that ISOPTIN (verapamil hydrochloride) tablets, 40 mg, 80 mg, and 120 mg, and CALAN (verapamil hydrochloride) tablets, 40 mg, 80 mg, 120 mg, and 160 mg, were withdrawn for reasons of safety or effectiveness. We have carefully reviewed our files for records concerning the withdrawal of ISOPTIN (verapamil hydrochloride) tablets, 40 mg, 80 mg, and 120 mg, and CALAN (verapamil hydrochloride) tablets, 40 mg, 80 mg, 120 mg, and 160 mg, from sale. We have also independently evaluated relevant literature and data for possible postmarketing adverse events. We have reviewed the available evidence and determined that these drug products were not withdrawn from sale for reasons of safety or effectiveness.

Accordingly, the Agency will continue to list ISOPTIN (verapamil hydrochloride) tablets, 40 mg, 80 mg, and 120 mg, and CALAN (verapamil hydrochloride) tablets, 40 mg, 80 mg, 120 mg, and 160 mg, in the “Discontinued Drug Product List” section of the Orange Book. The “Discontinued Drug Product List” delineates, among other items, drug products that have been discontinued from marketing for reasons other than safety or effectiveness. ANDAs that refer to ISOPTIN (verapamil hydrochloride) tablets, 40 mg, 80 mg, and 120 mg, and CALAN (verapamil hydrochloride) tablets, 40 mg, 80 mg, 120 mg, and 160 mg, may be approved by the Agency as long as they meet all other legal and regulatory requirements for the approval of ANDAs. If FDA determines that labeling for these drug products should be revised to meet current standards, the Agency will advise ANDA applicants to submit such labeling.

Dated: May 7, 2021.

Lauren K. Roth,

*Acting Principal Associate Commissioner for Policy.*